Docket No: AM101023 Application No: 10/792,280

Patent

IN THE SPECIFICATION

Applicants request that the Specification by amended by replacing the original numbered paragraphs with the following numbered marked-up replacement paragraphs.

[0024] FIG. 1 is a graphical representation showing that CAT2 and arginase type I (ARG1) are coinduced in Balb/c mice by both allergen (OVA) and recombinant murine IL-13 (IL-13). Briefly, Balb/c mice were sensitized to OVA by intraperitoneal injection on Day 0, challenged by intratracheal (IT) injection of either vehicle (phosphate buffered saline (PBS)) or OVA on Days 14 and 25, and the lungs harvested at Day 28. Additionally, nave Balb/c mice were treated IT with either PBS or IL-13 for 3 consecutive days and the lungs harvested at 72 hours. Total lung RNA was isolated and analyzed for mRNA expression using GENECHIP nucleic acid microarray GeneChip technology (AFFYMETRIX, Inc. Affymetrix) as described in Example 1. The mRNA frequency is expressed as parts per million.

[0025] FIG. 2 is a graphical representation showing that ARG1 expression is induced in Balb/c mice by both allergen (OVA) and recombinant murine IL-13 (IL-13). Briefly, Balb/c mice were sensitized to OVA by intraperitoneal injection on Day 0, challenged by intratracheal (IT) injection of either vehicle (phosphate buffered saline (PBS)) or OVA on Days 14 and 25, and the lungs harvested at Day 28. Additionally, naive Balb/c mice were treated IT with either PBS or IL-13 for 3 consecutive days and the lungs harvested at 72 hours. Total lung RNA was isolated and analyzed for mRNA expression using GENECHIP nucleic acid microarray GeneChip technology (AFFYMETRIX, Inc. Affymetrix) as described in Example 1. The mRNA frequency is expressed as parts per million.

[0033] FIG. 10 is a graphical representation showing that induction of ARG1 expression requires IL-4 receptor. Briefly, IL-4 receptor knockout mice (IL4R-/-) and IL-4 knockout mice (IL4-/-) sensitized to OVA, or treated with PBS or IL-13 as described in FIG. 1. Total lung RNA was isolated and analyzed for mRNA expression using <u>GENECHIP nucleic acid microarray GeneChip</u> technology (<u>AFFYMETRIX</u>, Inc. <u>Affymetrix</u>) as described in Example 1. The mRNA frequency is expressed as parts per million.

[0212] *In vitro* T7 polymerase driven transcription reactions for synthesis and biotin labeling of antisense cRNA, Qiagen Rneasy QIAGEN RNEASY RNA preparation material spin column

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purification and cRNA fragmentation were carried out as described (*supra*). GENECHIP GeneChip hybridization mixtures contained 10 μg fragmented cRNA, 0.5 mg/ml acetylated BSA, 0.1 mg/ml herring sperm DNA, in 1.times. MES buffer in a total volume of 200 μl as per manufacturer's instructions. Reaction mixtures were hybridized for 18 hr at 45 °C to AFFYMETRIX Affymetrix—Mu11KsubA and Mu11KsubB oligonucleotide arrays. The hybridization mixtures were removed and the arrays were washed and stained with Streptavidin R-phycoerythrin (Molecular Probes, Inc.) using the GeneChip GENECHIP Fluiditics Station 400 (AFFYMETRIX, Inc.Affymetrix) and scanned with a Hewlett Packard GENEARRAY GeneArray Scanner following Manufacture's instructions. Fluorescent data was collected and converted to gene specific difference averages using MicroArray Suite 4.0 software.